AMENDMENTS TO THE CLAIMS

Docket No.: 59526(47992)

- 1. (Previously presented) A method of inhibiting a viral infection in a subject, the method comprising administering to said subject a pharmaceutical composition comprising a ceramide-generating retinoid or a pharmaceutically acceptable salt thereof.
- 2. (Withdrawn) A method of preventing or inhibiting a viral infection in a subject, the method comprising administering to said subject a pharmaceutical composition comprising a ceramide-degradation inhibitor or a pharmaceutically acceptable salt thereof.
- 3. (Withdrawn) A method of preventing or inhibiting a viral infection in a subject, the method comprising administering to said subject a pharmaceutical composition comprising:
 - (a) a ceramide-generating retinoid or a pharmaceutically acceptable salt thereof; and
 - (b) a ceramide-degradation inhibitor or a pharmaceutically acceptable salt thereof.
- 4. (Original) A method of claim 1 wherein the ceramide-generating retinoid is a retinoic acid derivative.
- 5. (Withdrawn) A method of claim 1 wherein the ceramide degradation inhibitor is selected from the group consisting of glucosyl ceramide synthase inhibitors, sphingosine-1-phosphate synthesis inhibitors, protein kinase C inhibitors, and the pharmaceutically acceptable salts thereof.
- 6. (Previously presented) A method of inhibiting a viral infection, the method comprising administering a pharmaceutical composition comprising at least one N-(aryl)retinamide compounds to the subject suffering from or susceptible to a viral infection.
- 7. (Original) The method of claim 6, wherein the N-(aryl)retinamide modulates ceramide metabolism.
- 8. (Original) The method of claim 6, wherein the pharmaceutical composition comprises N-(4-hydroxyphenyl)retinamide or a derivative thereof.

9. (Original) The method of claim 6, wherein the pharmaceutical composition comprises at least one compound of the formula:

3

$$\bigcap_{\mathbb{R}^1} \bigcap_{(\mathbb{R}^2)_n} OH$$

wherein:

R¹ is hydrogen, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, or optionally substituted aralkyl;

R² is independently selected at each occurrence from the group consisting of hydrogen, halogen, hydroxy, optionally substituted alkoxy, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted amino, and optionally substituted mono- and di-alkylamino; and

n is an integer of from 0 to about 4.

- 10. (Original) The method of claim 6, wherein the pharmaceutical composition comprises N-(4-hydroxyphenyl)retinamide.
- 11. (Original) The method of claim 9, wherein the pharmaceutical composition further comprises one or more therapeutic agents selected from 1-phenyl-2-hexadecanoylamino-3-morpholino-1-propanol, chemokine inhibitors, HIV fusion inhibitors, viral protease inhibitors, reverse transcriptase inhibitors, and entry inhibitors.
 - 12. (Original) The method of claim 6, wherein the subject is a mammal.
 - 13. (Original) The method of claim 6, wherein the subject is a primate.
 - 14. (Original) The method of claim 6, wherein the subject is a human.

Docket No.: 59526(47992)

- 15. (Original) The method of claim 6, wherein the N-(aryl)retinamide compound inhibits HIV infectivity at a concentration of less than 10 μM.
- 16. (Original) The method of claim 6, wherein the N-(aryl)retinamide compound inhibits HIV infectivity at a concentration of less than 5µM.
- 17. (Original) A method of inhibiting HIV infectivity in a subject, the method comprising administration of N-(4-hydroxyphenyl)retinamide or a derivative thereof sufficient to increase ceramide levels in a cellular membrane susceptible to HIV entry.
- 18. (Original) The method of claim 17, wherein the N-(4-hydroxyphenyl)retinamide or a derivative thereof decreases the viral load in a subject by about 40%.
- 19. (Withdrawn) A method of inhibiting HIV infectivity in a subject, the method comprising administration of compound that stimulates the *de novo* synthesis of ceramide sufficient to increase ceramide levels in a cellular membrane susceptible to HIV entry.
- 20. (Withdrawn) The method of claim 19, wherein the compound that stimulates the generation of ceramide is sphingomyelinase.
- 21. (Withdrawn) The method of claim 19, wherein sphingomyelinase decreases viral load in a subject by about 40%, at least about 50%, 60%, 75%, 80%, 99.9%, up to about 100%.
- 22. (Withdrawn) The method of claim 21, wherein viral load is due to infection by HIV.
- 23. (Currently amended) A method of inhibiting a viral attachment/entry or exit phase of a virus by administering a pharmaceutical composition to a cell susceptible to infection by a virus, wherein the pharmaceutical composition comprises an inhibitor of at least one enzyme essential to ceramide metabolism, thereby inhibiting viral attachment/entry or exit phase of the virus.

- 24. (Original) The method of claim 23, wherein the enzyme is essential to a glycosylation step of ceramide metabolism.
- 25. (Original) The method of claim 23, wherein the pharmaceutical composition comprises at least one compound of the formula:

$$\bigcap_{\mathbb{R}^1} \bigcap_{(\mathbb{R}^2)_n} OH$$

Docket No.: 59526(47992)

wherein:

R¹ is hydrogen, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, or optionally substituted aralkyl;

R² is independently selected at each occurrence from the group consisting of hydrogen, halogen, hydroxy, optionally substituted alkoxy, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted amino, and optionally substituted mono- and di-alkylamino; and

n is an integer of from 0 to about 4.

- 26. (Original) The method of claim 25, wherein the pharmaceutical composition comprises N-(4-hydroxyphenyl)retinamide.
- 27. (Original) The method of claim 26, wherein the pharmaceutical composition further comprises one or more therapeutic agents selected from 1-phenyl-2-hexadecanoylamino-3-morpholino-1-propanol, chemokine inhibitors, HIV fusion inhibitors, viral protease inhibitors, reverse transcriptase inhibitors, and entry inhibitors.
 - 28. (Original) The method of claim 25, wherein the cell is a mammalian cell.
 - 29. (Original) The method of claim 28, wherein the cell is an immune cell.
 - 30. (Previously presented) The method of claim 25, wherein the virus is HIV.

Docket No.: 59526(47992)

- 31. (Original) The method of claim 25, wherein the N-(aryl)retinamide compound inhibits HIV infectivity at a concentration of less than 10 μ M.
- 32. (Original) The method of claim 25, wherein the N-(aryl)retinamide compound inhibits HIV infectivity at a concentration of less than $5\mu M$.
- 33. (Previously presented) The method of claim 25, wherein the at least one enzyme essential to ceramide metabolism is sphingomyelinase and the inhibitor inhibits the viral attachment/entry phase of the virus in a cell by about 40% up to about 100%.
 - 34. (Withdrawn) A kit comprising:
 - a) one or more agents for increasing ceramide concentration of a cell,
 - b) means for detecting at least one of a) ceramide concentration of the cells, and 2) inhibition of viral infectivity of the cell; and
 - c) directions for using the kit.
- 35. (Withdrawn) The kit of claim 34, wherein the agents comprise a pharmaceutical composition of a N-aryl retinamide compound capable of activating ceramide biosynthesis in addition to a pharmaceutical composition that inhibits ceramide glycosolation and (glyco)sphingolipid formation.
- 36. (Withdrawn) The kit of claim 34, wherein the agents comprise any one or more of compositions as identified by Formula I and substituted groups thereof.
 - 37. (New) The method of claim 28, wherein the cell is a differentiated macrophage.